

H36.D2.B7 ANTI-TISSUE FACTOR LIGHT CHAIN VARIABLE REGION

GACATTCAGATGACCCAGTCTCCTGCCTCCCAGTCTGCATCTCTGGGAGAAAGTGCACCATCACATGC
D I Q M T Q S P A S Q S A S L G E S V T I T C

CTGGCAAGTCAGACCATTGATACATGGTTAGCATGGTATCAGCAGAAACCAGGAAATCTCCTCAGCTC
L A S Q T I D T W L A W Y Q Q K P G K S P Q L

CTGATTTATGCTGCCACCAACTTGGCAGATGGGGTCCCATCAAGTTTCAGTGGCAGTGGATCTGGCACA
L I Y A A T N L A D G V P S R F S G S G S G T

AAATTTCTTCAAGATCAGCAGCCCTACAGGCTGAAGATTTTGTAATTATTACTGTCAACAAGTTTAC
K F S F K I S S L Q A E D F V N Y Y C Q Q V Y

AGTTCTCCATTACGTTTCGGTGCTGGGACCAAGCTGGAGCTGAAA
S S P F T F G A G T K L E L K

* CDR REGIONS UNDERLINED.

FIG. 1A

H36.D2.B7 ANTI-TISSUE FACTOR HEAVY CHAIN VARIABLE REGION

GAGATCCAGCTGCAGCAGTCTGGACCTGAGCTGGTGAAGCCTGGGGCTTCAGTGCAGGTATCCTGCAAG
 E I Q L Q Q S G P E L V K P G A S V Q V S C K
 ACTTCTGGTTACTCATTCACTGACTACAACGTGTACTGGTGAGGCAGAGCCCATGGAAAGAGCCTTGAG
 T S G Y S F T D Y N V Y
 TGGATTGGATATATTGATCCTTACAATGGTATTACTATCTACGACCAGAACTTCAAGGGCAAGGCCACA
 W I G Y I D P Y N G I T I Y D Q N F K G K A T
 TTGACTGTTGACAAAGTCTTCCACCACAGCCTTCATGTCATCTCAACAGCCTGACATCTGACGACTCTGCA
 L T V D K S S T T A F M H L N S L T S D D S A
 GTTTATTCTGTGCAAGAGATGTGACTACGGCCCTTGACTTCTGTGGGCCAAGGCACCACTCTCACAGTC
 V Y F C A R D V T T A L D F
 TCCTCA
 S S

* CDR REGIONS UNDERLINED.

FIG. 1B

ANTIBODY	APPARENT K_d M ⁻¹	APPARENT K_d M
BY ELISA		
D2	5.2×10^9	1.9×10^{-10}
I47	6.5×10^9	1.5×10^{-10}
K73	9.8×10^9	1.0×10^{-10}
K80	2.3×10^9	4.3×10^{-10}
L102	2.5×10^9	4.0×10^{-10}
L133	1.7×10^9	5.9×10^{-10}
BY BIACore		
H36	3.1×10^{10}	3.2×10^{-11}
I43	2.3×10^9	4.3×10^{-10}
I47	3.2×10^9	3.1×10^{-10}
L133	4.6×10^9	2.2×10^{-10}
M107	1.1×10^9	9.1×10^{-10}

FIG. 2

ANTIBODY NAME	% INHIBITION ANTIBODY PREINCUBATED WITH TF/VIIa
D1	0
D1B	1
H31	4
<u>H36</u>	<u>95</u>
I43	1
J131	7
K80	0
K82	0
K87	1
L97B	7
L101	0
L102	0
L105	0
L133	0
M5	1
M107	34

FIG. 3

ANTIBODY NAME	<u>% INHIBITION</u> TF PREINCUBATED WITH ANTIBODY PRIOR TO ADDITION OF VIIa	<u>% INHIBITION</u> TF PREINCUBATED WITH VIIa PRIOR TO ADDITION OF ANTIBODY
D1	15	nd
D1B	48	12.7
H31	64	21
H36	0	0
I43	68	55
J131	38	11
K80	12	nd
K82	0	nd
K87	0	nd
L96	0	nd
L101	38	11
L102	14	nd
L105	4	nd
L133	13	nd
M5	0	nd
M107	0	nd

FIG. 4

[rhTF],nM	[H36.D2],nM	H36.D2/rhTF MOLAR RATIO	CLOTTING TIME (SECONDS)	% INHIBITION OF rhTF FUNCTION
0.0048	0	0	102.3	0
	1.61	335.4	114.3	31.3
	3.23	670.8	121.3	45.8
0.023	0	0	77.6	0
	1.61	70.0	85.3	52.2
	3.23	140.0	91.1	65.2
	6.45	280.4	99.6	73.9
0.092	0	0	49.3	0
	3.23	35.1	65.8	65.2
	6.45	70.1	88.5	90.2
	12.90	140.2	113.3	95.7
0.46	0	0	32.6	0
	6.45	14.0	52.7	82.4
	12.90	28.0	80.2	96.7
	32.30	70.2	117.9	99.3
2.30	0	0	23.9	0
	16.10	7.0	47.1	94.4
	32.30	14.0	95.2	99.7
	64.50	28.0	115.3	99.9
11.52	0	0	22.2	0
	16.10	1.4	30.2	93.4
	32.30	2.8	46.0	98.8
	64.50	5.6	87.6	99.9
	161.30	14.0	114.0	100.0

FIG. 5

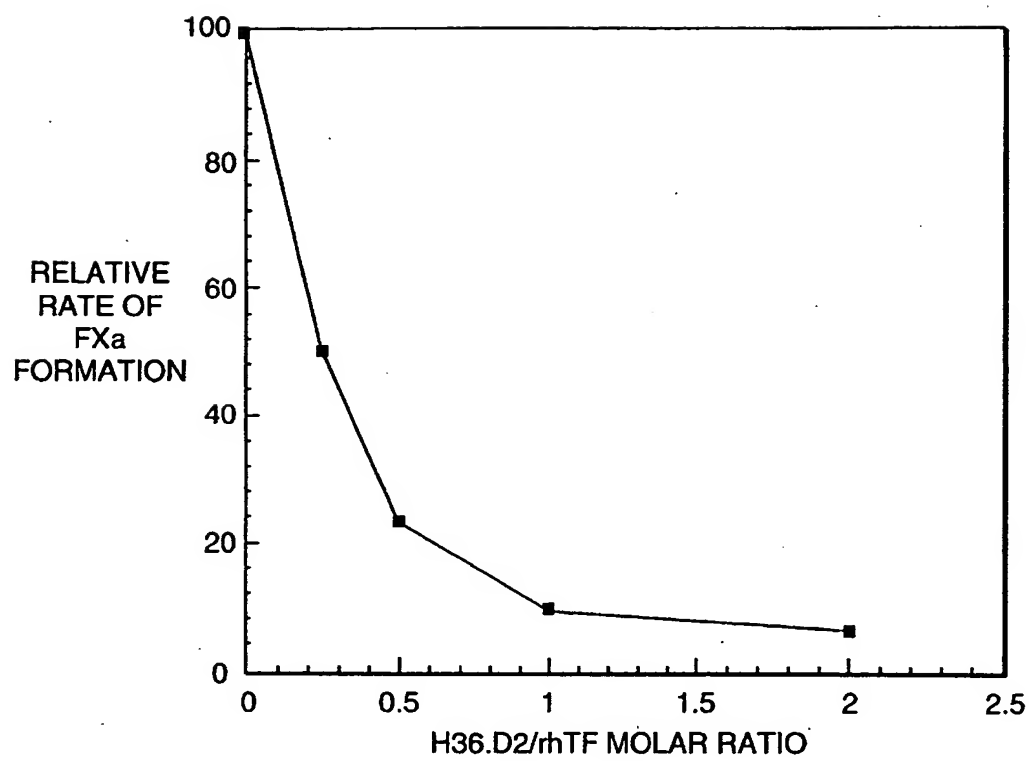


FIG. 6A

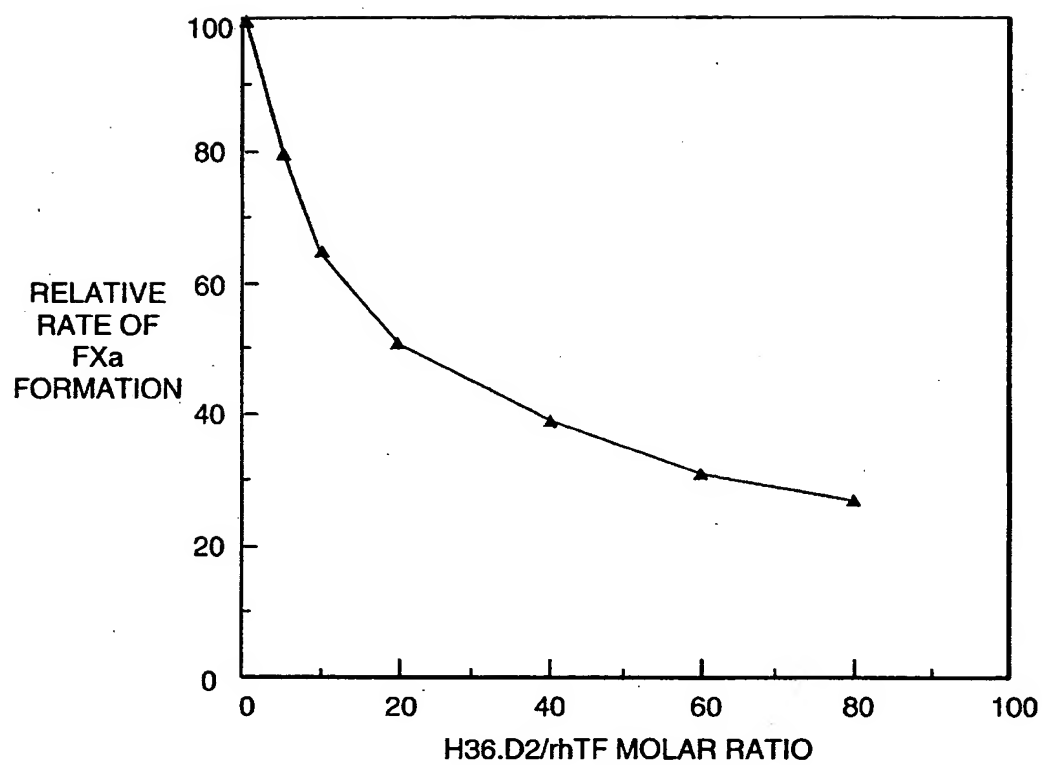


FIG. 6B

H36.D2 CONCENTRATION (ng)	<u>% INHIBITION</u> CELLS (TF/FVII) AND H36.D2 PREINCUBATED PRIOR TO FX ADDITION	<u>% INHIBITION</u> FX AND H36.D2 ARE ADDED SIMULTANEOUSLY TO CELLS (TF/FVII)
0	0	0
50	88	nd
100	92	nd
200	97	nd
800	nd	76
1600	nd	78
3200	nd	92

FIG. 7

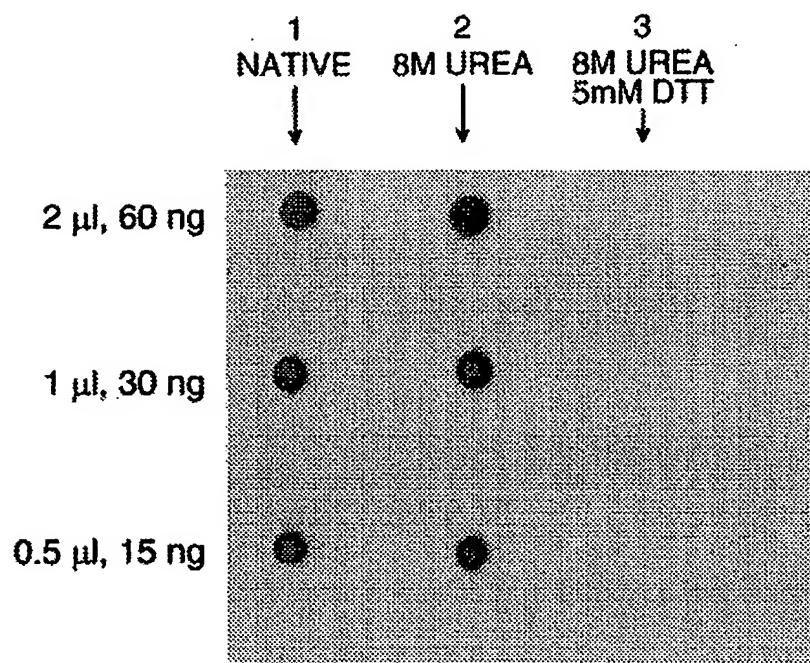


FIG. 8A

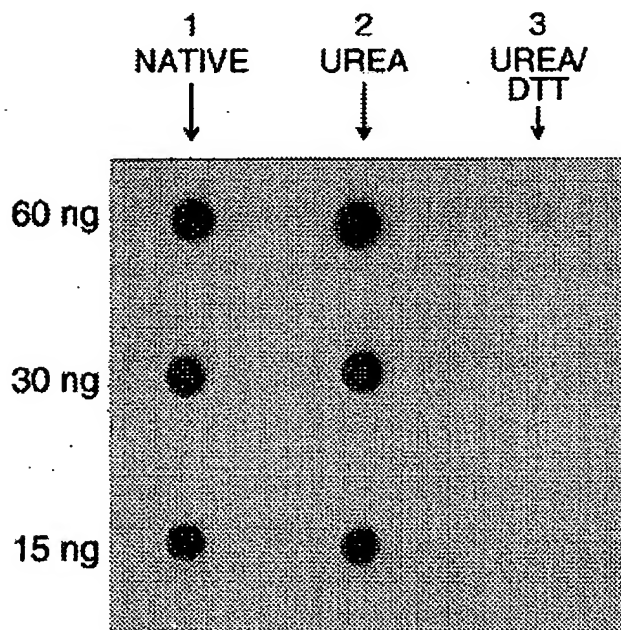


FIG. 8B